

FIG. 1A

1 CCCCCGCTCGGTCTTCCACCTCACCTTTCGAGCTGGCCGCCGCTTGCTGTGCGCAGTTTC 60
61 GGGGGACTGGACCTTCCCTGGCTTTTAGCAGCGCCGAGCGCCATGGCGACCTTTTGCTGG 120
121 GCAGGTGACCGATTCCGGGTGCCGAAGGAGCTGGCGTGGGTCTGCCTTGCAGCCGCCCG 180
181 CCTGGACAGGATGTTTGCTAGAGGGCTGAAGAGGAAATATGGTGACCAGGAAGAAGGAGT 240
1 M F A R G L K R K Y G D Q E E G V 17
241 AGAGGGTTTGGCACTGTCCCTTCTATAGCCTGCAGCGACAGTCACTCCTGGACATGTC 300
E G F G T V P S Y S L Q R Q S L L D M S 37
301 CCTTGTCAAGCTCCAGCTCTGTACATGCTAGTGGAGCCCAATCTCTGCCGCTCGGTCTT 360
L V K L Q L C H M L V E P N L C R S V L 57
361 CATCGCCAACACAGTCCGGCAGATCCAGGAGGAAATGAGCCAGGATGGTGTGTGGCATGG 420
I A N T V R Q I Q E E M S Q D G V W H G 77
421 GATGGCACCCAGAAATGTAGATCGGGCACCAGTTGAACGCCTGGTGTCCACAGAGATCCT 480
M A P Q N V D R A P V E R L V S T E I L 97
481 GTGTCGTACAGTGAGGGGAGCTGAGGAAGAGCACCTGCTCCTGAACCTGGAAGATGCTCC 540
C R T V R G A E E E H P A P E L E D A P 117
541 CTTGCAAACTCGGTTTCCGAGCTCCCATCGTTGGCTCAGCACCAGGGCAAAGGAACCC 600
L Q N S V S E L P I V G S A P G Q R N P 137
601 TCAGAGCAGCCTCTGGGAGATGGACAGCCCAAGAAACAGGGGAAGCTTTCAGAAGTC 660
Q S S L W E M D S P Q E N R G S F Q K S 157
661 ACTGGACCAGATATTTGAGACCTTGAGAACAAAACTCCAGTTCAGTGGAGGAACCTTT 720
L D Q I F E T L E N K N S S S V E E L F 177
721 CTCAGATGTGGACAGCTCCTACTATGACCTGGACACAGTGCTAACAGGAATGATGAGTGG 780
S D V D S S Y Y D L D T V L T G M M S G 197
781 GACCAAGTCCAGTCTCTGCAATGGCCTTGAGGGCTTTGCTGCAGCCACCCCTCCTCCCAG 840
T K S S L C N G L E G F A A A T P P P S 217
841 TTCCACTTGCAAGTCTGACCTGGCTGAGCTGGACCATGTGGTAGAGATTCTGGTGGAGAC 900
S T C K S D L A E L D H V V E I L V E T 237
901 CTGAGAGGCCACCCAGTGGGCTAAGGGTGAGGCCACCAGTCCCATGGAGCTCACGTGT 960
*
961 GTTGTGACCCAGAGACAGATAAGCACTTGTCTTAAGAGGGGCTCTGGCTCTTGAGCTCAT 1020
1021 TATCCTTTTGTGTGACATTGGACTCACTGTGGAGGATGGTGTGTACAGCTATGTCTAGT 1080
1081 CTATTTTCAATTAGATAGGTGAACCTTTCTAAAATTAAGTTTTATATGTTTTTGGGCAATA 1140
1141 TTTTGTCTTAAGATATATTTTTTAAACTTTTTATACTTTAGATTTTTTTTCAGCTATTTTC 1200
1201 TTAAGAGTATATTTTTTCTACAAACATCCTCTGCTGCTACATTAGAAACATTTATAACCT 1260
1261 AAATACGATTGGTGTGTGTCATTTTAAAGGTTTAAATAGAAAACCTCTTTTGTACTGAGTC 1320
1321 TCTACACTCCCAAGGCAACTGTAAATGTAGCCGGCCGGGTGTTTACATGAGAGGCTCCAG 1380
1381 TATGGTCTACATTCTAGTAGAGCTTGAAAAGAACCATGCACAGCTCCACTGCCCCCTCAC 1440
1441 TGGGTCTGCTCTGGCGGATCGGAGCTCTCTTCTAGCCCCGTGTGCAGGATGGCTTTATT 1500
1501 TATGCCTATTTATATGTAAATGCCACTGAAAGCTAAGGTCTTACTCCTGGAAATCCCAAC 1560
1561 ACCAGTTCTTCAGGGACTGCTGTGAGGCAGTGCCTTATGCAGGTCTTGTCTTGGCCATC 1620
1621 ACTGTCTGGTTCCAGCCAGCACATGTGACATGAGGACATGACATGCCGAACCACCCA 1680
1681 GCACCACATGCTCCATGTCAAGTGTGTACGTGGAGACCAGTGGCTCCAGGCCTGTGCTC 1740
1741 AGAGAGGGTGTGCAGTCTTACGTGTGCTGGGGGGGACGACGGTGACCTGTGCTTGTGCT 1800
1801 TTTTAAATGGTGTCTTGGACGTTTTAAGGTTAAAAACAATCCGACTCCATATGATTTAGG 1860
1861 GCTCCTCCACCTGGGGTGGCCCTATGCTGTCTGCTTGGATCTCAAAGTCTTGGTACTC 1920
1921 GGCAGTGTGAGTCCACCCCATGTATCCTTTTTGTTTCTTGTGCTTTTTTTGGACTT 1980
1981 CCAACCTGAGCCTAAGGTTTTATTTATATGTGCTTCAATATCAACAATGTAAACCTCA 2040
2041 CTTTATTAAGTATCCAGCAATGGAAAAAAAAAAAAAAAAA

FIG. 1B

1 GGGGAAGCTGGCGGCACAGCCGTGGCGCCTGGCTGAGCAGAGGACCCGGCGGGCGGCCTCG 60
61 CGGGTCAGGACACAATGTTTGCACGAGGACTGAAGAGGAAATGTGTTGGCCACGAGGAAG 120
1 M F A R G L K R K C V G H E E D 16
121 ACGTGGAGGGAGCCCTGGCCGGCTTGAAGACAGTGTCTCATACAGCCTGCAGCGGCAGT 180
V E G A L A G L K T V S S Y S L Q R Q S 36
181 CGCTCCTGGACATGTCTCTGGTGAAGTTGCAGCTTTGCCACATGCTTGTGGAGCCCAACC 240
L L D M S L V K L Q L C H M L V E P N L 56
241 TGTGCCGCTCAGTCCTCATTGCCAACACGGTCCGGCAGATCCAAGAGGAGATGACGCAGG 300
C R S V L I A N T V R Q I Q E E M T Q D 76
301 ATGGGACGTGGCGCACAGTGGCACCCAGGCTGCAGAGCGGGCGCCGCTCGACCGCTTGG 360
G T W R T V A P Q A A E R A P L D R L V 96
361 TCTCCACGGAGATCCTGTGCCGTGCAGCGTGGGGGCAAGAGGGGGCACATCCTGCTCCTG 420
S T E I L C R A A W G Q E G A H P A P G 116
421 GCTTGGGGGACGGCCACACAGGGTCCAGTTTCTGACCTTTGCCCAGTCACCTCAGCAC 480
L G D G H T Q G P V S D L C P V T S A Q 136
481 AGGCACCAAGGCACCTGCAGAGCAGCGCCTGGGAGATGGATGGCCCTCGAGAAAACAGAG 540
A P R H L Q S S A W E M D G P R E N R G 156
541 GAAGCTTTTACAAGTCACTTGATCAGATATTTGAAACGCTGGAGACTAAAAACCCAGCT 600
S F H K S L D Q I F E T L E T K N P S C 176
601 GCATGGAAGAGCTGTTCTCAGACGTGGACAGCCCTACTACGACCTGGACACAGTACTGA 660
M E E L F S D V D S P Y Y D L D T V L T 196
661 CAGGCATGATGGGGGGTGCCAGGCCGGGCCCTGCGAAGGGCTCGAGGGCTTGGCTCCGG 720
G M M G G A R P G P C E G L E G L A P A 216
721 CCACCCAGGCCCTAGCTCCAGCTGCAAGTCCGACCTGGGCGAGCTGGACCAGTGGTGG 780
T P G P S S S C K S D L G E L D H V V E 236
781 AGATCCTGGTGGAGACCTGAGCAGGAGCCCTGAGTGCTCACAGCCGCCTCTGACGCATTG 840
I L V E T * 241
841 ACACGTGAGCACTGGCTCCCACGGAGGGTGCGCCTGCCGCCAGCGGCCAGCCTTGCTGC 900
901 CCTGTCTGCTGATTCTGAGAAATCCCAGAACAGCCATTACCAGTGGGGCTGCAGCCCTA 960
961 GGCCCGTCCCCTCACCTCCCCCTGTGGAGCGCCAGGCAGAGGCTGTTCTGGAAGGCTT 1020
1021 CTTGTCTTCTGACGTCCCCACAGCCCTGGGCCCTCGTGTCTCTTTGTGTCCCCCACTGT 1080
1081 AGAGGACGGTGAGCCGCAGCTGCATCAACCTCCTTTTACCTTTAGATAGGTGAATTTTA 1140
1141 CAATTCAGTTTTACATGTTTTGGGCAGTATTTTGTCTTAAGATATATTTTTTAACTTTT 1200
1201 TATACCTTATCTCTTTAGATTTTTTTCAGCTATTTTCTTAAAGTATATTTTTTCTATAAA 1260
1261 CATCCTTTGCTGCTACATTAGAATTTTATAGCCTAAACAATTGCAGTTGGTGTGTTTCA 1320
1321 TTTTTTTAAGGTTTAAATAAGGGTTTTTGTGTTTGTGTTTTTGTGTTTTTGCAGTGAGCATCAC 1380
1381 TACAGTCTCAGTCAACAGTGTGAATGTATCATGTTTTACTTTAAATGTGTGTGTGATACT 1440
1441 TCTTCATTATGTCTGCTGCGCTGCAGTGAGACCTGGGTGAAAATCAGGAGCCGCACACAGCC 1500
1501 ACATCTTCTAGACCTAAGAGTAAATTATGGAGGATTTTATTTATGTCTATTTATATGTA 1560
1561 AATGTCATTGAAGACAAAGGTCAAATATTTGTCTGTTGTAGATCACAGGCACCAAGTTGG 1620
1621 TCTTCAGGGACCTCATAGCCCTCGGTGGTGCCTTCTCAAGGCAGTGTCTCGGAGGCTC 1680
1681 CCATCAGGGTCAGCCCATGCACCTGCCCTGGGTGAGGAAGTAGCATTGCTGCTGGATGAG 1740
1741 AAACGCCTGCGCTGCTCTGTTAGACTGGTGCTGAAACAAAAGGTTAAGGCTAGGTTGAAG 1800
1801 TCTAGAATGAAAGAAATCTGAATCCATGTCATTATAACCCCTTGATCTGTAGTGTGATG 1860
1861 GGTGCTGCCGCAGGCAGGGAGTGAGCTGGGGGTGCCTGCAGCCTTCCACTCCTGCCCCGC 1920
1921 CTCACCCACATGCTCCCTGTTTCTCATGCTTTCTCTAACTTCCTCACCCTTAACCAAA 1980
1981 AAGGTGTGTTTTCTTTTGTGCATATAGCCATTCTTAAATATCAGTGATGTAAACCTCACT 2040
2041 TTATTAAAAAATTATCCAGCAAAAAAAAAAAAAAAAAAAAAA

FIG. 2

Mouse	Hepp	1	MFARGLKRKYG	---	DOEEGV	EGFG	TVPS	YS	LQ	RQ	SLLD	MSLV	KLQ	LCH	ML	VE	PN	LC	RSV																																			
Human	HEPP	1	MFARGLKRKC	CV	GHEED	VE	GAL	AG	LK	TVSS	YS	LQ	RQ	SLLD	MSLV	KLQ	LCH	ML	VE	PN	LC	RSV																																
Mouse	Hepp	57	LIANTVRQ	IQEEMS	QDGV	WHG	MAPQ	NVD	RAP	VER	LV	STE	ILC	RTV	RGA	EE	EH	PA	PE	LE	DA																																	
Human	HEPP	61	LIANTVRQ	IQEEMT	QDGI	WRT	VAPQ	AAE	RAP	LD	RL	V	STE	ILC	RAA	WQ	EG	AHP	AP	GL	GDG																																	
Mouse	Hepp	117	PLQNSV	SELP	IV	GS	APG	Q	RNP	QSS	LW	EMD	SPQ	ENR	GSF	QK	S	L	DQ	I	F	E	T	L	E	N	K	N	S	S	V	E	E	L																				
Human	HEPP	121	HTQGPV	SDLC	PV	TS	AQ	A	PH	LQ	S	S	A	W	EMD	G	P	R	E	N	R	G	S	F	H	K	S	L	D	Q	I	F	E	T	L	E	T	K	N	P	S	C	M	E	E	L								
Mouse	Hepp	177	FSDVD	SSYY	D	L	T	V	L	T	G	M	M	S	G	T	K	S	S	L	C	N	G	L	E	G	F	A	A	T	P	P	P	S	S	T	C	K	S	D	L	A	E	L	D	H	V	V	E	I	L	V	E	
Human	HEPP	181	FSDVD	SPYY	D	L	T	V	L	T	G	M	M	G	G	A	R	P	G	P	C	E	G	L	E	G	L	A	P	A	T	P	G	P	S	S	S	C	K	S	D	L	G	E	L	D	H	V	V	E	I	L	V	E
Mouse	Hepp	237	T																																																			
Human	HEPP	241	T																																																			

FIG. 3

Zebrafish Hepp

Mouse Hepp

Human HEPP

1	MFSKGTIKRKFA	DGGEEISDDGLVAARVASS	YSLQRQSLLDMSLKLKLQ	CHMLVEPNLCRS
1	MFARGLKRKYG	---DQEEGVGEFGTV	PSYSLQRQSLLDMSLVKLQ	CHMLVEPNLCRS
1	MFARGLKRKCVGH	-EEDVEGALLAGLKT	VSSYSLQRQSLLDMSLVKLQ	CHMLVEPNLCRS

Zebrafish Hepp

Mouse Hepp

Human HEPP

61	VLIANTVRQIQEEMTHD	GSWHMVT	EAF	CGASQSP	SERLVATEVL	CR
56	VLIANTVRQIQEEMSQ	DDGVWHGM	APQNVDR	-APVERLVSTEIL	CR	TVRGAE
60	VLIANTVRQIQEEMTQ	DTMRTVAPQAAER	-APLDR	LVSTEIL	CR	AAWGQEG
						GAHPAPGL

FIG. 4B

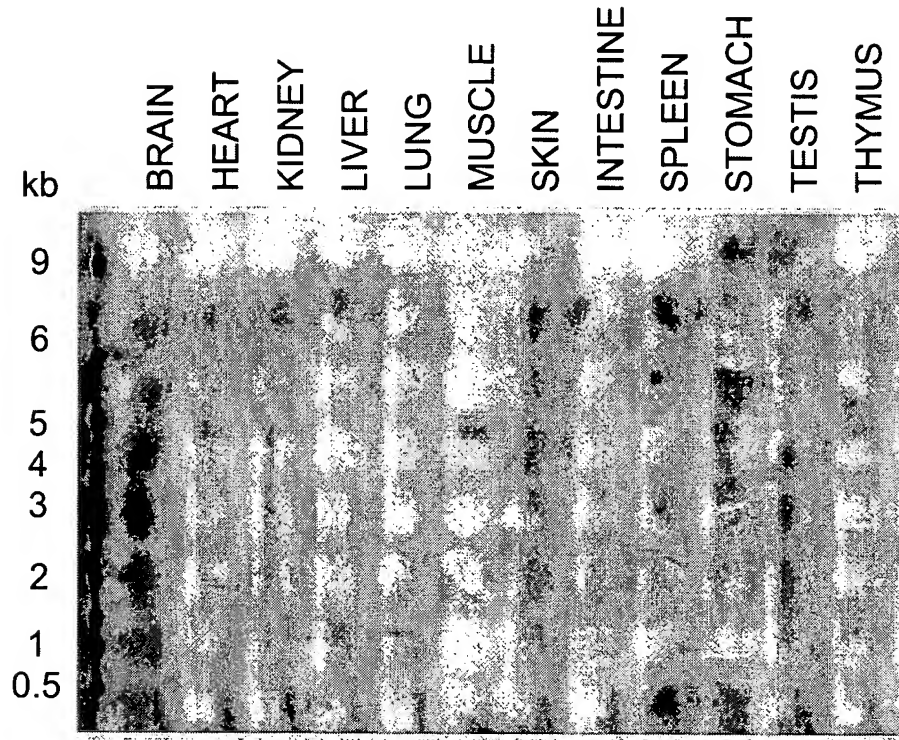


FIG. 4A

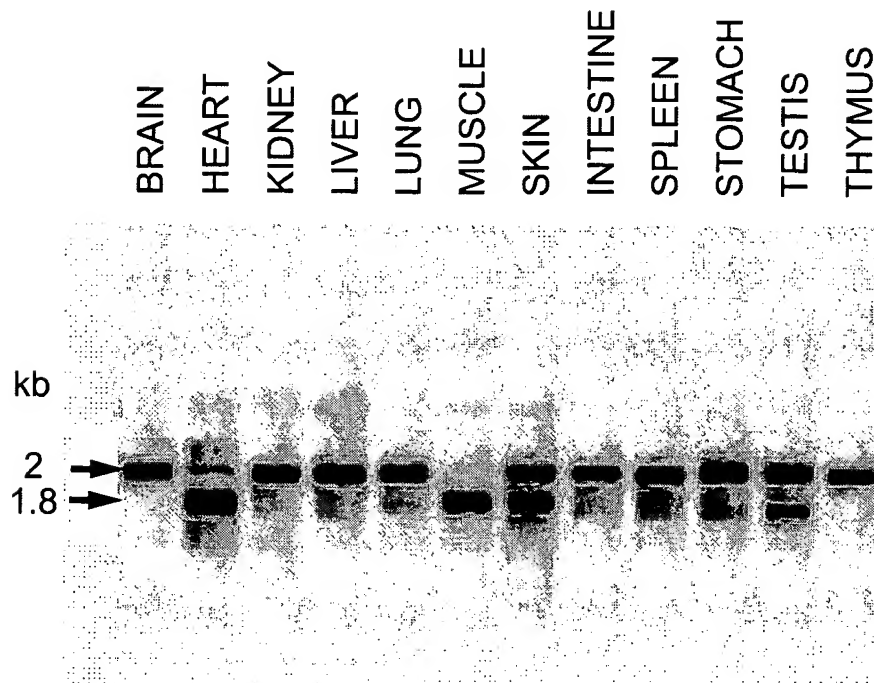


FIG. 5

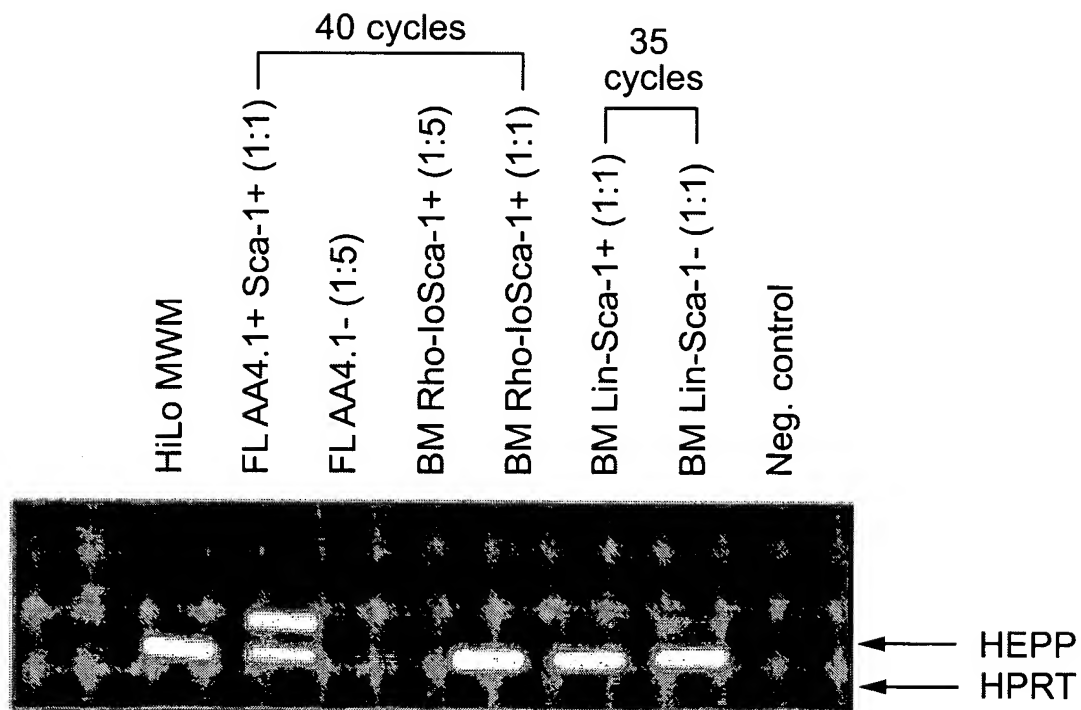


FIG. 6A

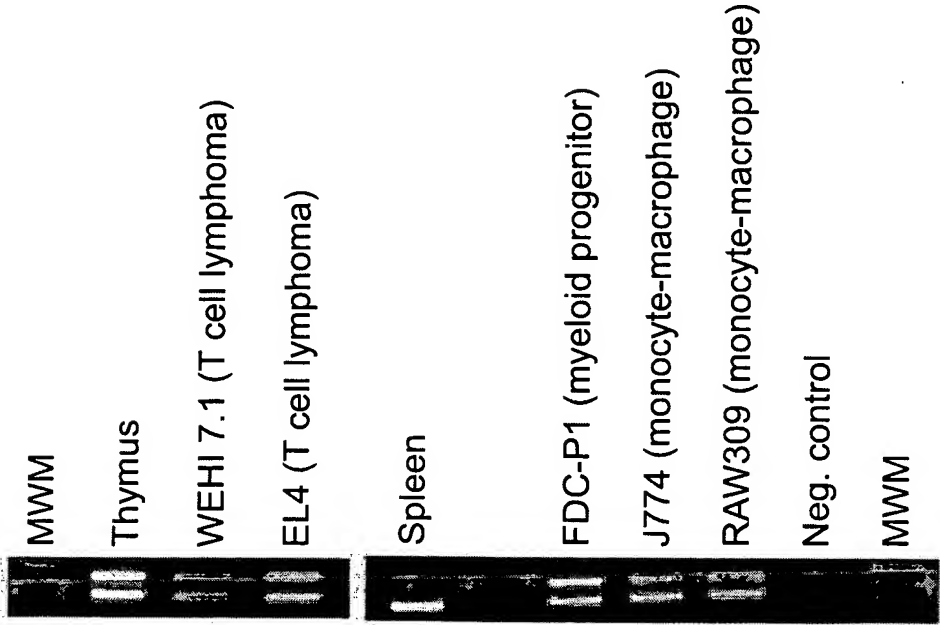


FIG. 6B

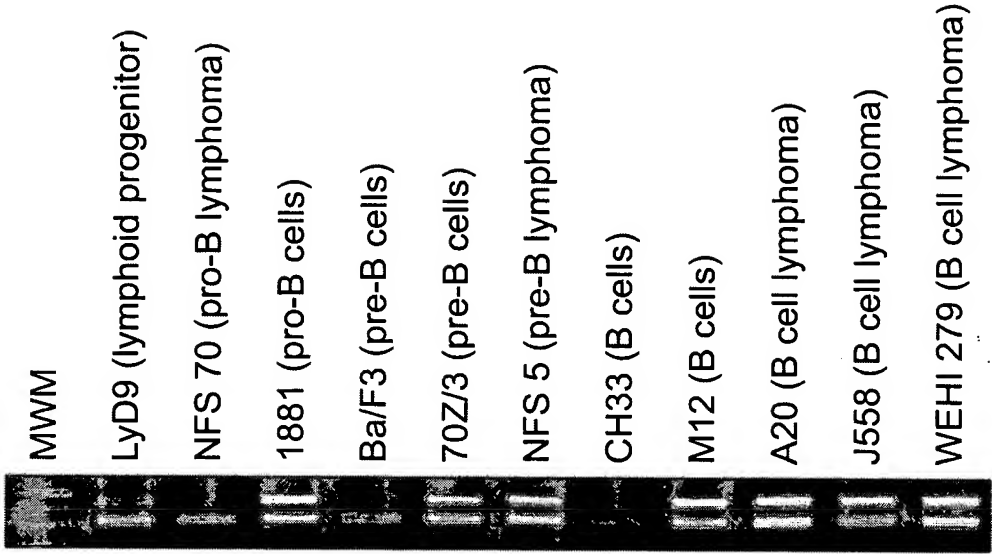
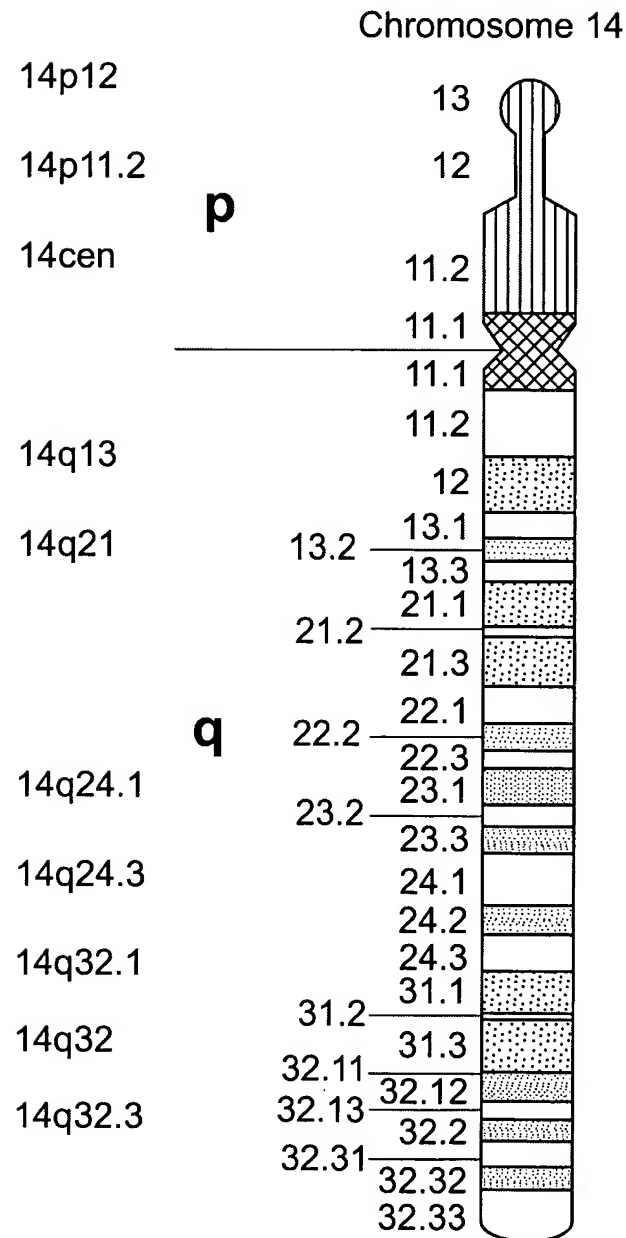


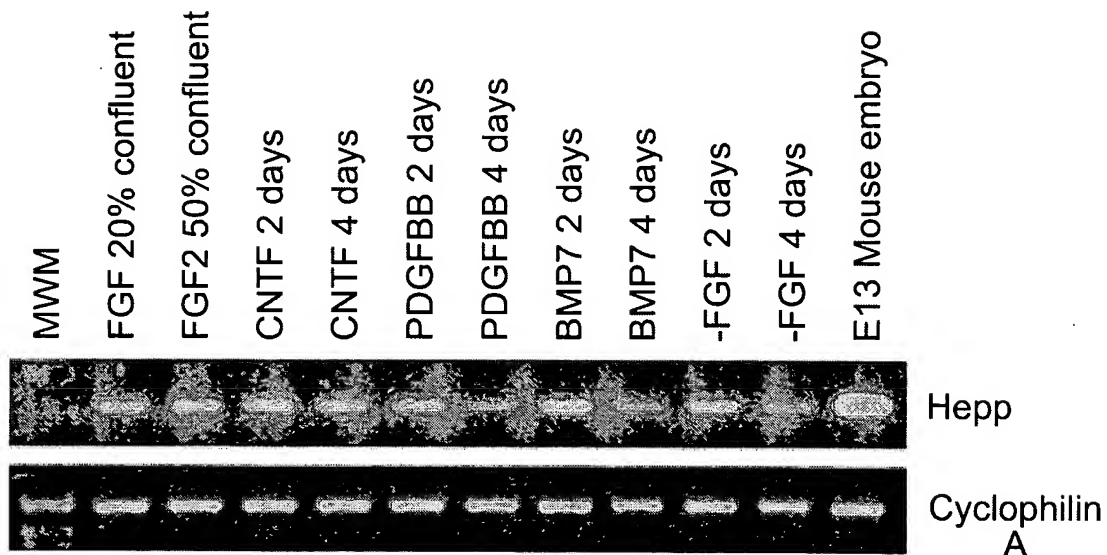
FIG. 7



translocation breakpoints

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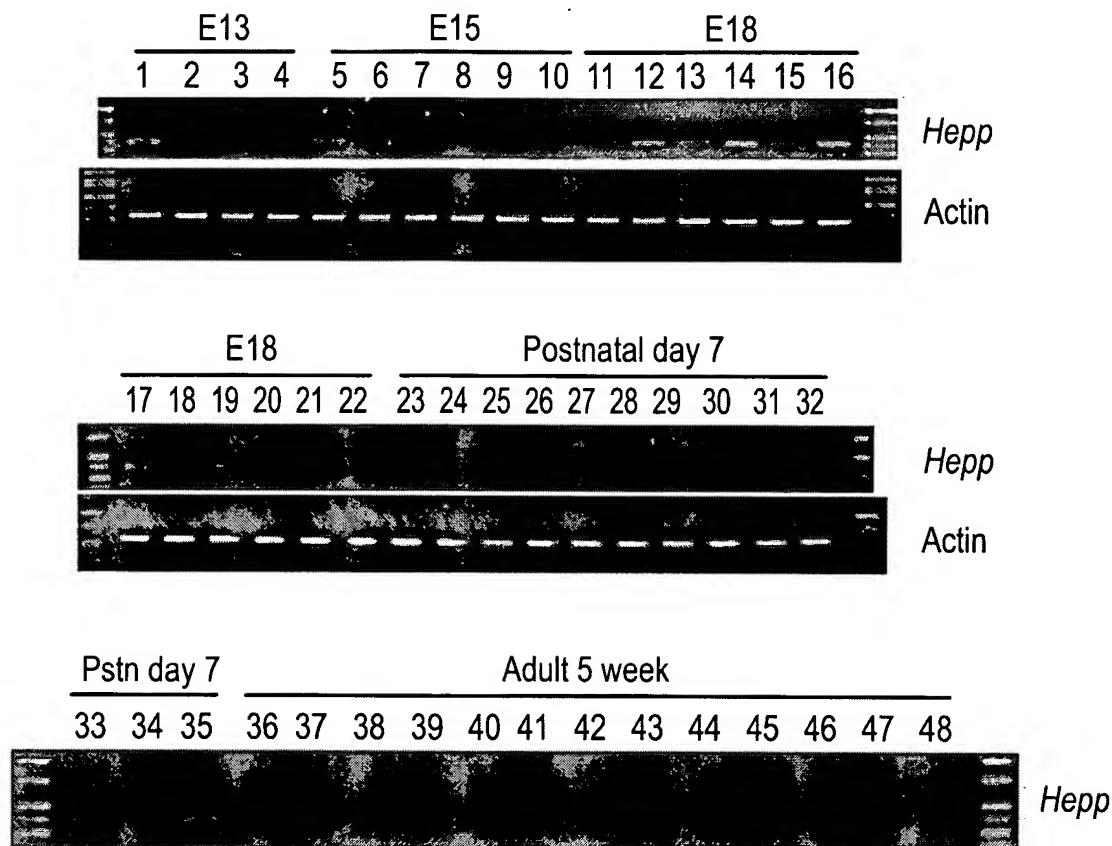
FIG. 8



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FIG. 9



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Embryo day 13

1. Teleocephalon/Diencephalon
2. Mesococephalon (Midbrain)
3. Rhombococephalon (Hindbrain)
4. Spinal cord

Embryo day 15

5. Teleocephalon
6. Diencephalon
7. Midbrain
8. Pons
9. Medulla
10. Spinal cord

Embryo day 18

11. Frontal cortex
12. Posterior cortex
13. Entorhinal cortex
14. Olfactory bulb
15. Hippocampus
16. Striatum
17. Thalamus
18. Hypothalamus
19. Midbrain
20. Pons
21. Medulla
22. Spinal cord

Postnatal day 7

23. Frontal cortex
24. Posterior cortex
25. Entorhinal cortex
26. Olfactory bulb
27. Hippocampus
28. Striatum
29. Thalamus
30. Hypothalamus
31. Cerebellum
32. Midbrain
33. Pons
34. Medulla
35. Spinal cord

Adult 5 week

36. Frontal cortex
37. Posterior cortex
38. Entorhinal cortex
39. Olfactory bulb
40. Hippocampus
41. Striatum
42. Thalamus
43. Hypothalamus
44. Cerebellum
45. Midbrain
46. Pons
47. Medulla
48. Spinal cord

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FIG. 10A

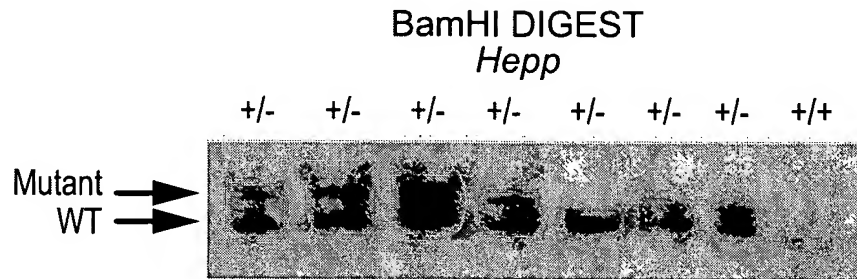


FIG. 10B

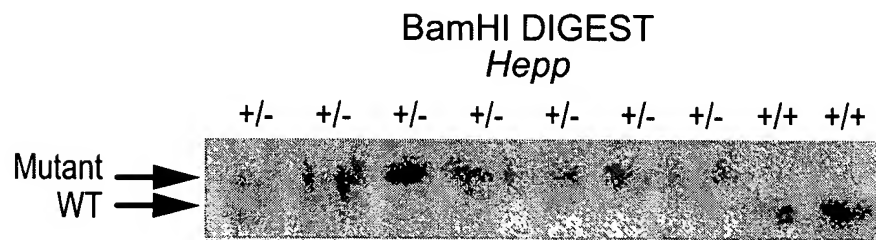
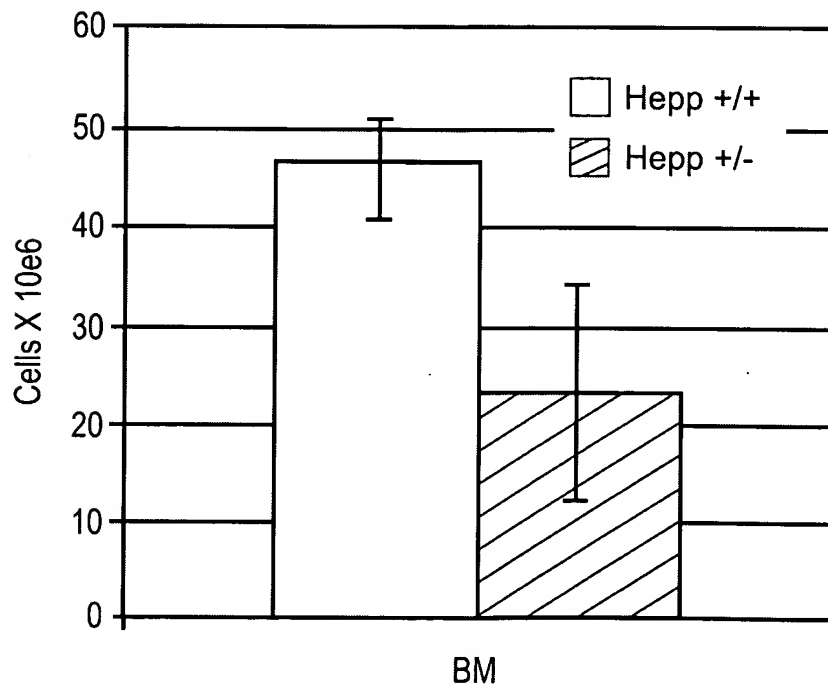


FIG. 11



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FIG. 12

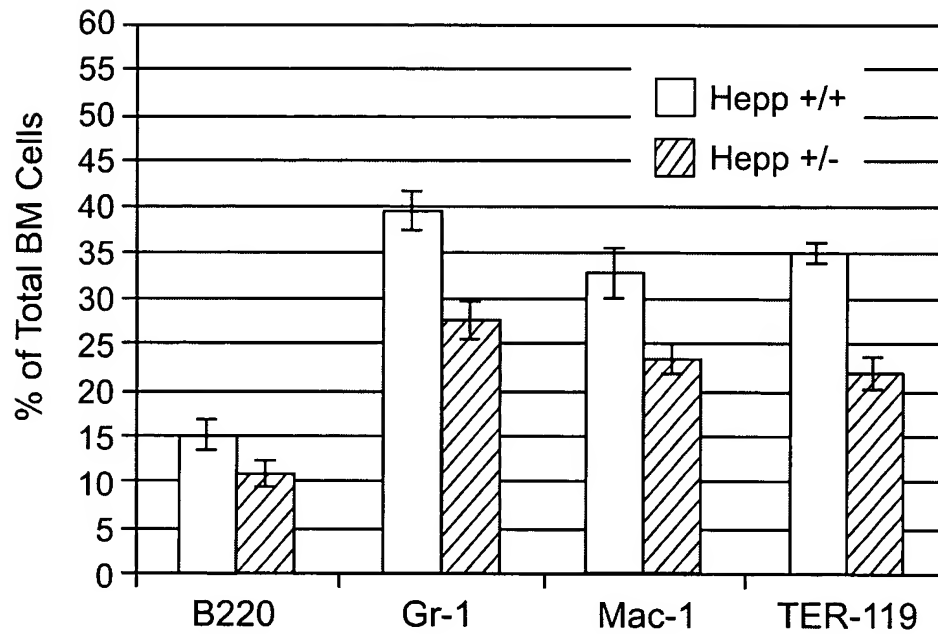
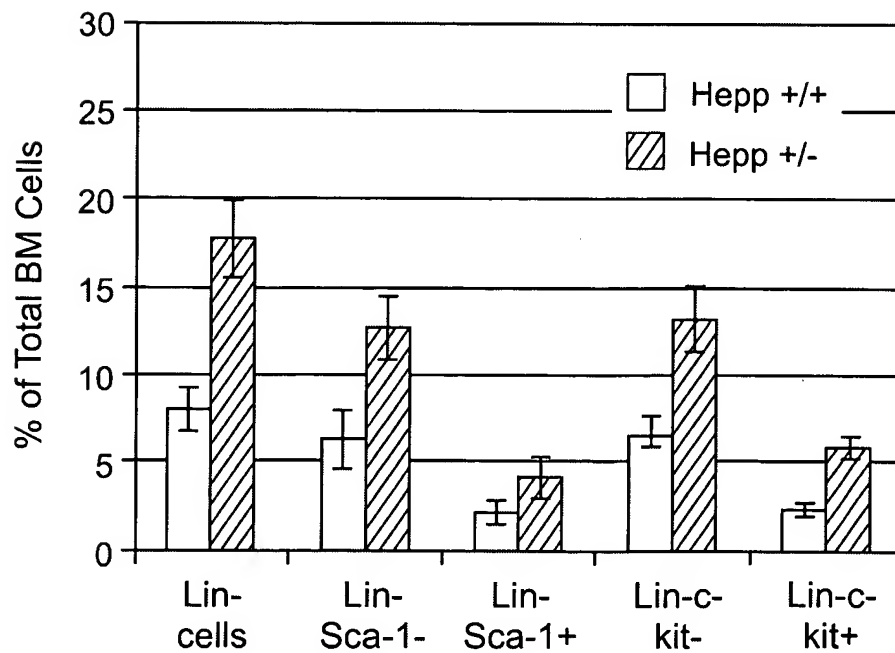
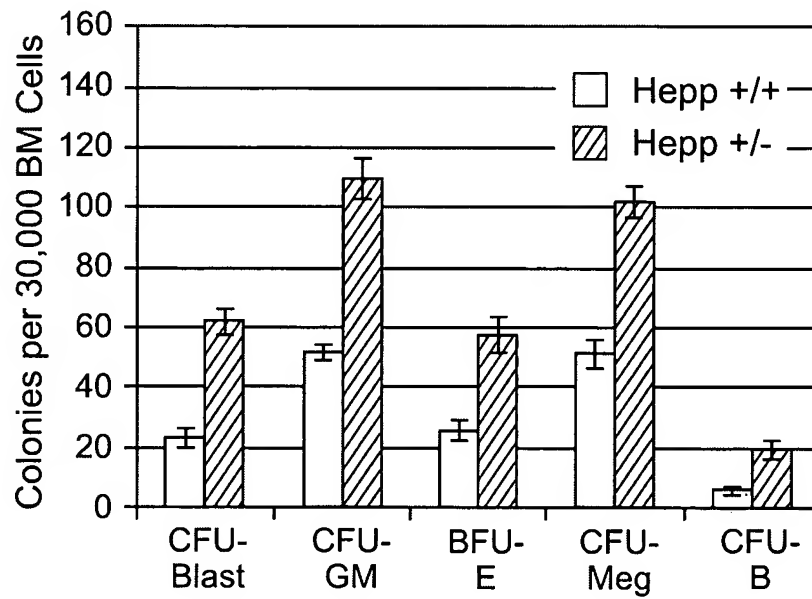


FIG. 13



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FIG. 14



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FIG. 15A

Hepp +/+

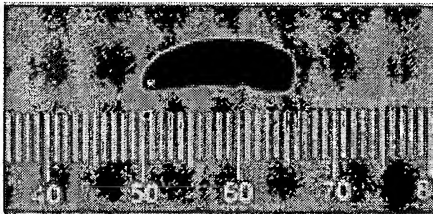


FIG. 15B

Hepp +/-

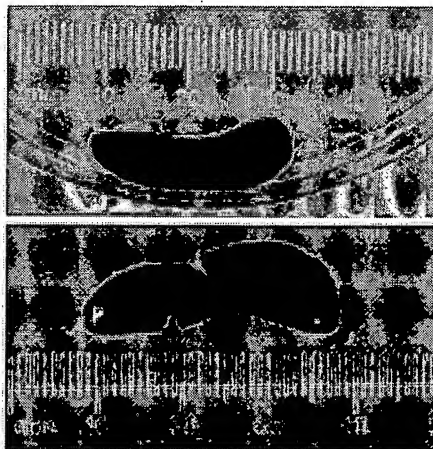
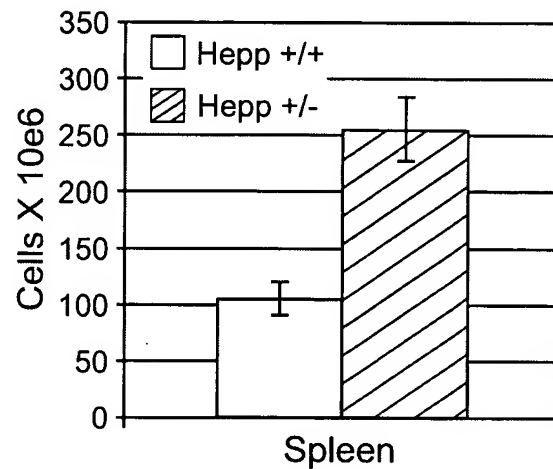


FIG. 15C



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FIG. 16A

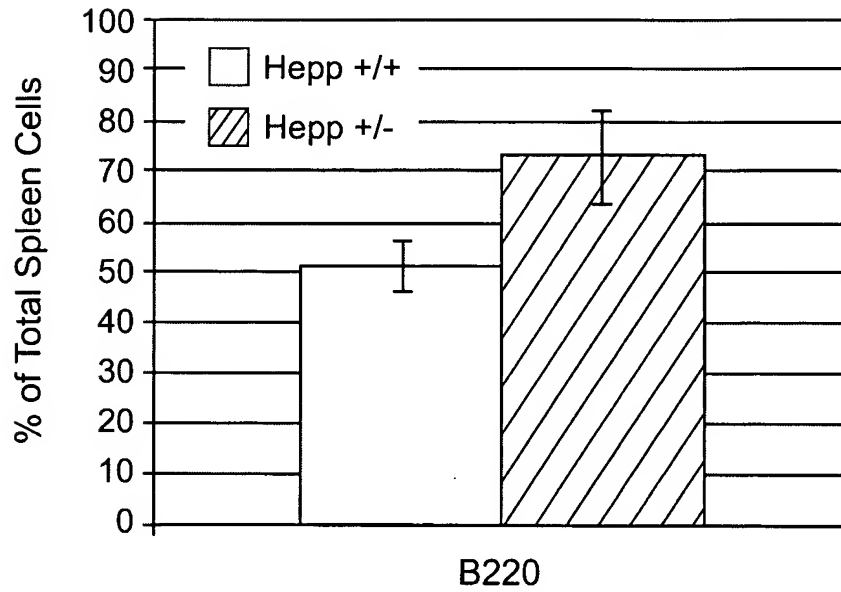


FIG. 16B

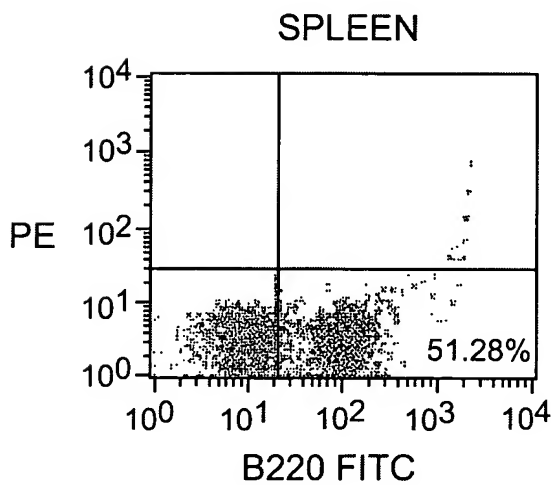
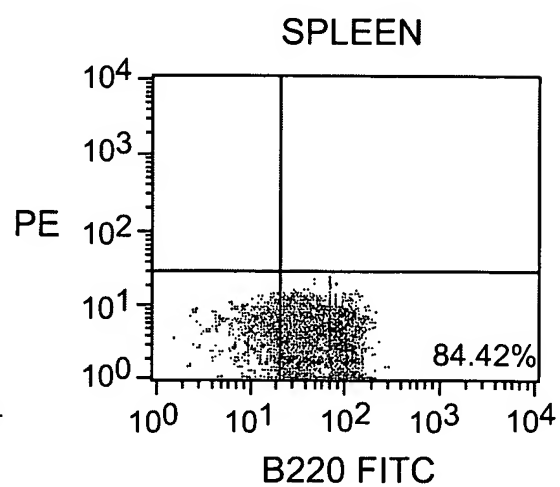


FIG. 16C



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FIG. 17A

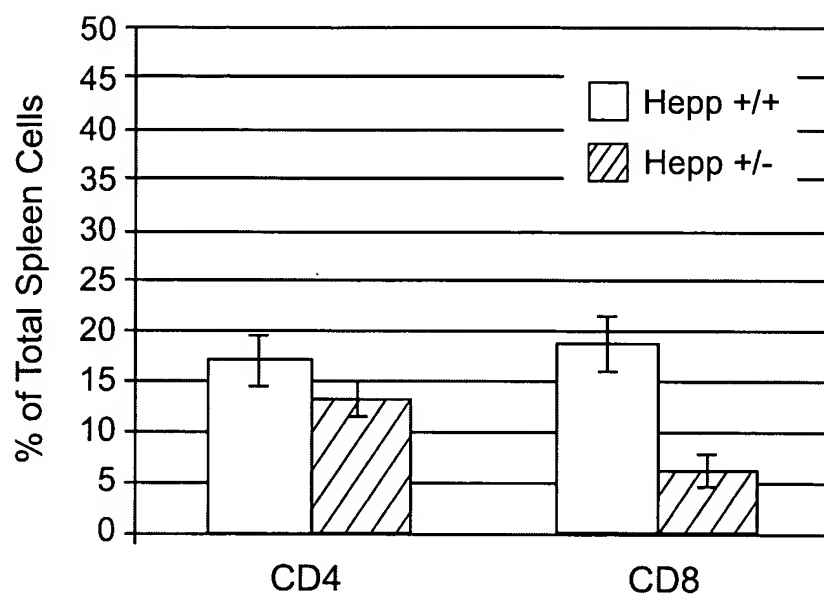


FIG. 17B

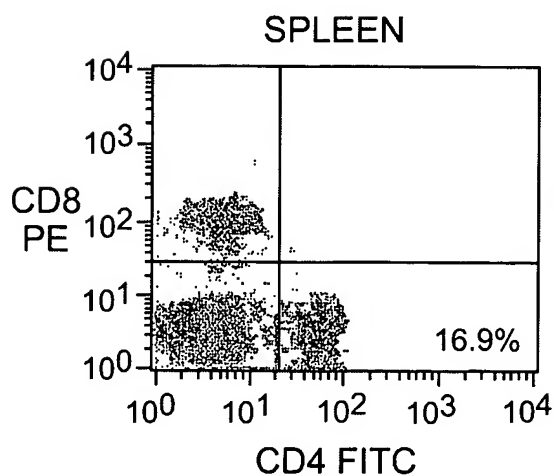


FIG. 17C

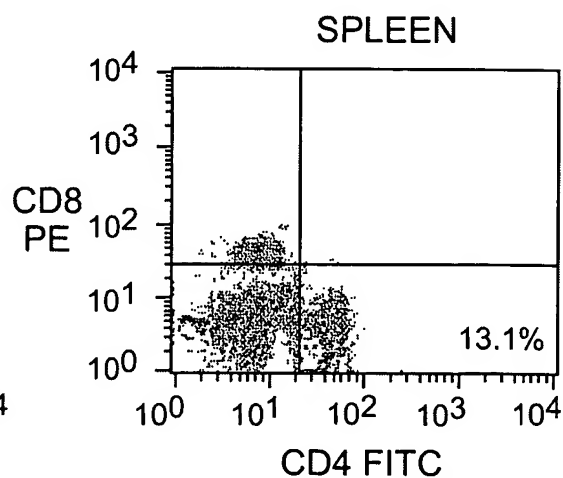


FIG. 18A

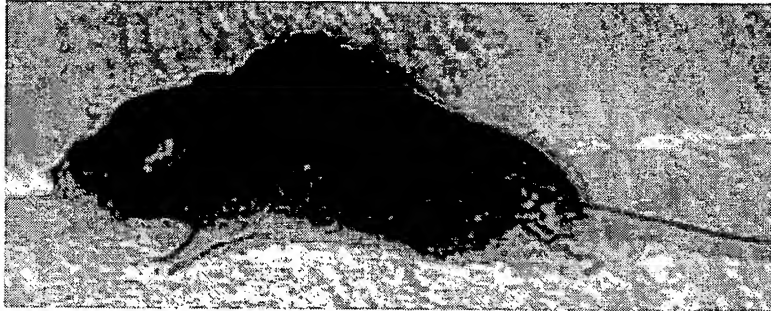


FIG. 18B

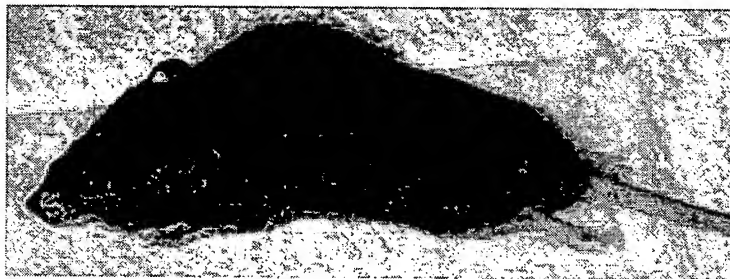
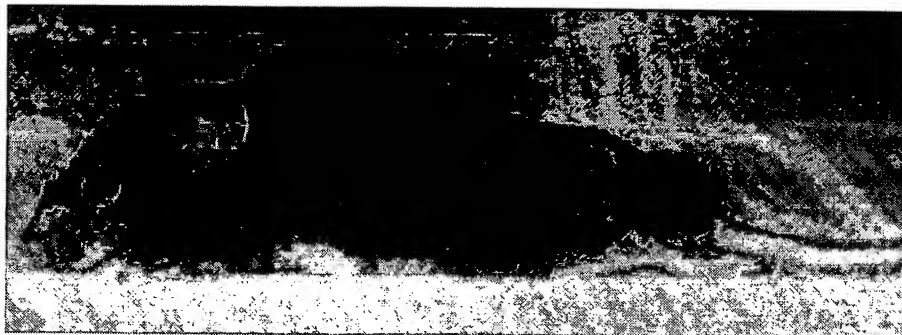


FIG. 18C



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